## SEQUENCE LISTING

(1)	GENERAL INFORMATION:
(i)	APPLICANT: GREENE, ET AL.
(ii)	TITLE OF INVENTION: Human Tumor Necrosis Factor
	Receptor
(iii)	NUMBER OF SEQUENCES: 9
	A DEPENDE
(iv)	CORRESPONDENCE ADDRESS:
	(A) ADDRESSEE: CARELLA, BYRNE, BAIN, GILFILLAN,
	(A) ADDRESSEE: CARELLA, BYRNE, BAIN, GILFILLAN, CECCHI, STEWART & OLSTEIN
	(B) STREET: 6 BECKER FARM ROAD
	(C) CITY: ROSELAND
	(D) STATE: NEW JERSEY
	(E) COUNTRY: USA
sub)	(F) ZIP: 07068
0:/	(2) 222
(v)	COMPUTER READABLE FORM:
	(A) MEDIUM TYPE: 3.5 INCH DISKETTE
	(B) COMPUTER: IBM PS/2
	(C) OPERATING SYSTEM: MS-DOS
	(D) SOFTWARE: WORD PERFECT 5.1
(vi)	CURRENT APPLICATION DATA:
	(A) APPLICATION NUMBER:
	(B) FILING DATE: Concurrently
	(C) CLASSIFICATION:
	\ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \
(vii)	PRIOR APPLICATION DATA
	(A) APPLICATION NUMBER: PCT/US95/03216
	(B) FILING DATE: 15 MAR 1995

(viii	) ATTORNEY/AGENT INFORMATION:	
	(A) NAME: FERRARO, GREGORY D.	
	(B) REGISTRATION NUMBER: 36,134	
	(C) REFERENCE/DOCKET NUMBER: 325800-381	
	(3)	
( ÷ \	TELECOMMUNICATION INFORMATION:	
(ix)	(A) TELEPHONE: 201-994-1700	
	(B) TELEFAX: 201-994-1744	
•	GEO TE NO 1	
(2)	INFORMATION FOR SEQ ID NO:1:	
(i)	SEQUENCE CHARACTER STICS	
	(A) LENGTH: 1173 BASE PAIRS	
	(B) TYPE: NUCLEIC ACLD	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: cDNA	
(22)		
(vi)	SEQUENCE DESCRIPTION: SEQ ID NO:1:	
(XI)	bligolines substitution and substitution	
አጥር አ አ (	CAAGT TGCTGTGCTG CGCGCTCGTG TTTCTGGACA TCTCCATTAA GTGGACCACC	60
CAGGA	AACGT TTCCTCCAAA GTACCTTCAT TATGACGAAG AAACCTCTCA TCAGCTGTTG	120
	CAAAT GTCCTCCTGG TACCTACCTA AAACAACACT GTACAGCAAA GTGGAAGACC	180
GTGTG	CGCCC CTTGCCCTGA CCACTACTAC ACAGACAGCT GGCACACOAG TGACGAGTGT	240
CTATA	CTGCA GCCCCGTGTG CAAGGAGCTG CAGTACGTCA AGCAGGAGTG CAATCGCACC	300
	CCGCG TGTGCGAATG CAAGGAAGGG CGCTACCTTG AGATAGAGTT CTGCTTGAAA	360 420
	GAGCT GCCCTCCTGG ATTTGGAGTG GTGCAAGCTG GAACCCCAGA GCGAAATACA	480
	CAAAA GATGTCCAGA TGGGTTCTTC TCAAATGAGA CGTCATCTAA AGCACCCTGT ACACA CAAATTGCAG TGTCTTTGGT CTCCTGCTAA CTCAGAAAGG AAATGCAACA	540
	CAACA TATGTTCCGG AAACAGTGAA TCAACTCAAA AATGTGGAAT AGATGTTACC	600
	TGAGG AGGCATTCTT CAGGTTTGCT GTTCCTACAA AGTTTACGCC TAACTGCTT	660
	CTTGG TAGACAATTT GCCTGGCACC AAAGTAAACG CAGAGAGTGT AGAGAGATA	720
	GCAAC ACAGCTCACA AGAACAGACT TTCCAGCTGC TGAAGTTATG GAAACATCAA	780
	AGACC AAGATATAGT CAAGAAGATC ATCCAAGATA TTGACCTCTG TGAAAACAGC	840
	GCGGC ACATTGGACA TGCTAACCTC ACCTTCGAGC AGCTTCGTAG CTTGATGGAA	900

960

4050

AGCTTACCGG GAAAGAAGT GGGAGCAGAA GACATTGAAA AAACAATAAA GGCATGCAAA

CCCAGTGACC AGATCCTGAA GCTGCTCAGT TTGTGGCGAA TAAAAAATGG CGACCAAGAC

ACCTTGAAGG GCCTAATGCA CGCACTAAAG CACTCAAAGA CGTACCACTT TCCCACAAAC 1080
TGTCACTGAG AGTCTAAAGA AGACCATCAG GTTCCTTCAC AGCTTCACAA TGTACAAATT 1140
GTATCAGAAG TTATTTTTAG AAATGATAGG TAA 1173

- (2) INFORMATION FOR SEQ ID NO:2:
  - (i) SEQUENCE CHARACTERISTICS
    - (A) LENGTH: 390 AMINO ACIDS
    - (B) TYPE: AMINO ACID
    - (C) STRANDEDNESS:
    - (D) TOPOLOGY: LINEAR
  - (ii) MOLECULE TYPE: PROTEIN
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Met Asn Lys Leu Leu Cys Cys Ala Leu Val Phe Leu Asp Ile Ser
-20 -15 -10

Ile Lys Trp Thr Thr Gln Glu Thr Phe Pro Pro Lys Tyr Leu His

-5 1 5
Tyr Asp Glu Glu Thr Ser His Gln\Leu Leu Cys Asp Lys Cys Pro

10 15 20

Pro Gly Thr Tyr Leu Lys Gln His Cys Thr Ala Lys Trp Lys Thr 25 30 35

Val Cys Ala Pro Cys Pro Asp His Tyr Tyr Thr Asp Ser Trp His

Thr Ser Asp Glu Cys Leu Tyr Cys Ser Pro Val Cys Lys Glu Leu 55

Gln Tyr Val Lys Gln Glu Cys Asn Arg Thr His Asn Arg Val Cys
70 75 80

Glu Cys Lys Glu Gly Arg Tyr Leu Glu Ile Glu Phe Cys Leu Lys

His Arg Ser Cys Pro Pro Gly Phe Gly Val Val Gln Ala Gly Thr

Pro Glu Arg Asn Thr Val Cys Lys Arg Cys Pro Asp Gly Phe Phe
115 120 125

Ser Asn Glu Thr Ser Ser Lys Ala Pro Cys Arg Lys His Thr Asn Cys Sex Val Phe Gly Leu Leu Leu Thr Gln Lys Gly Asn Ala Thr His Asp Ask Ile Cys Ser Gly Asn Ser Glu Ser Thr Gln Lys Cys Gly Ile Asp Val Thr Leu Cys Glu Glu Ala Phe Phe Arg Phe Ala Val Pro Thr Lys Phe Thr Pro Asn Trp Leu Ser Val Leu Val Asp Asn Leu Pro Gly Thr Lys Val Asn Ala Glu Ser Val Glu Arg Ile Lys Arg Gln His Ser Ser Gln Glu Gln Thr Phe Gln Leu Leu Lys Leu Trp Lys His Gln Asn Lys Asp Gln Asp Ile Val Lys Lys Ile Ile Gln Asp Ile Asp Leu Cys Glu Asn Ser Val Gln Arg His Ile Gly His Ala Asn Leu Thr Phe Glu Gln Leu Arg Ser Leu Met Glu Ser Leu Pro Gly Lys Lys Val Gly Alà Glu Asp Ile Glu Lys Thr Ile Lys Ala Cys Lys Pro Ser Asp Gln Ite Leu Lys Leu Leu Ser Thr Leu Lys Gly Leu Leu Trp Arg Ile Lys Asn Gly Asp Gln Asp Met His Ala Leu Lys His Ser Lys Thr Tyr Hig Phe Pro Thr Asn Cys His Ser Glu Ser Lys Glu Asp His Gln Val Pro Ser Gln Leu His Asn Val Gln Ile Val Ser Glu Val Ile Phe Arg\Asn Asp Arg 

## (2) INFORMATION FOR SEQ ID NO:3:

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(i)	SEQUENCE CHARACTERISTICS	
	(A) LENGTH: 33 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:3:	
GCC	AGAGGAT CCGAAACGTT TCCTCCAAAG TAC	33
(2)	INFORMATION FOR SEQ ID NO:4:	
	$\alpha$	
(i)	SEQUENCE CHARACTERISTICS	
	(A) LENGTH: 33 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID \	
	(C) STRANDEDNESS: SINGLE \	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO 4:	
CGG	CTTCTAG AATTACCTAT CATTTCTAAA AAT	33
(2)	INFORMATION FOR SEQ ID NO:5:	
(i)	SEQUENCE CHARACTERISTICS	
	(A) LENGTH: 31 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide \	

(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:5:	
GCGC	GGATCC ATGAACAAGT TGCTGTGCTG C	31
(2)	INFORMATION FOR SEQ ID NO:6:	
(i)	SEQUENCE CHARACTERISTICS	
	(A) LENGTH: 34 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:6:	
GCGC	CTCTAGA TTACCTATCA TTTCTAAAAA TAAC	34
(2)	INFORMATION FOR SEQ ID NO.7:	
(i)	SEQUENCE CHARACTERISTICS	
	(A) LENGTH: 31 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:7:	
GCG	CGGTACC TCAGTGGTTT GGGCTCCTCC C	31
(2)	INFORMATION FOR SEQ ID NO:8:	
(i)	SEQUENCE CHARACTERISTICS	

	(A) LENGTH: 39 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:8:	
GCCAG	AGGAT CCGCCACCAT GAACAAGTTG CTGTGCTGC	39
(2)	INFORMATION FOR SEQ NO:9:	
(i)	SEQUENCE CHARACTERISTICS	
• •	(A) LENGTH: 60 BASE PAIRS	
	(B) TYPE: NUCLEIC ACID	
	(C) STRANDEDNESS: SINGLE	
	(D) TOPOLOGY: LINEAR	
(ii)	MOLECULE TYPE: Oligonucleotide	
(xi)	SEQUENCE DESCRIPTION: SEQ ID NO:9	
CGGCTI	CTAG AATCAAGCGT AGTCTGGGAC GTCGTATGGG TACCTATCAT TTCTAAAAAT	60
		60

- A process for producing a polypeptide comprising: expressing from the host cell of Claim 5 the polypeptide encoded by said DNA.
- 7. A process for producing cells capable of expressing a polypeptide comprising transforming or transfecting the cells with the vector of Claim 4.
- 8. A receptor polypeptide comprising a member selected from the group consisting of:
- (i) a polypeptide having the deduced amino acid sequence of SEQ ID NO:2 and fragments, analogs and derivatives thereof; and
- (ii) a polypeptide encoded by the cDNA of ATCC Deposit No. 75899 and fragments, analogs and derivatives of said polypeptide.
- 9. An antibody against the polypeptide of claim &
- 10. A compound which activates the polypeptide of claim 8.
- 11. A compound which inhibits activation the polypeptide of claim 8
- 12. A method for the treatment of a patient having need to activate a TNF receptor comprising: administering to the patient a therapeutically effective amount of the compound of claim 10.
- 13. A method for the treatment of a patient having need to inhibit a TNF receptor comprising: administering to the patient a therapeutically effective amount of the compound of claim 11.

- 14. The method of claim 12 wherein said compound is a polypeptide and a therapeutically effective amount of the compound is administered by providing to the patient DNA encoding said agonist and expressing said agonist *in vivo*.
- The method of claim 13 wherein said compound is a polypeptide and a therapeutically effective amount of the compound is administered by providing to the patient DNA encoding said antagonist and expressing said antagonist in vivo.
- 16. A method for identifying compounds which bind to and activate the receptor polypeptide of claim 8 comprising:

contacting a cell expressing on the surface thereof the receptor polypeptide, said receptor being associated with a second component capable of providing a detectable signal in response to the binding of a compound to said receptor polypeptide, with a compound under conditions sufficient to permit binding of the compound to the receptor polypeptide; and

identifying if the compound is capable of receptor binding by detecting the signal produced by said second component.

17. A method for identifying compounds which bind to and inhibit activation of the polypeptide of claim 8 comprising:

thereof the receptor polypeptide, said receptor being associated with a second component capable of providing a detectable signal in response to the binding of a compound to said receptor polypeptide, with an analytically detectable ligand known to bind to the receptor polypeptide

and a compound to be screened under conditions to permit binding to the receptor polypeptide; and

determining whether the compound inhibits activation of the polypeptide by detecting the absence of a signal generated from the interaction of the ligand with the polypeptide.

18. A process for diagnosing a disease or a susceptibility to a disease related to an under-expression of the polypeptide of claim & comprising:

determining a mutation in the nucleic acid sequence encoding said polypeptide.

- 19. The polypeptide of Claim & wherein the polypeptide is a soluble fragment of the polypeptide and is capable of binding a ligand for the receptor.
- 20. A diagnostic process comprising:
  analyzing for the presence of the polypeptide of
  claim 19 in a sample derived from a host.

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